Process for the preparation of heterologous bispecific antibodies

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The invention pertains to a method for producing heterologous bispecific antibodies. The object of the invention is a method designed to yield heterologous bispecific antibodies with a high level of purity. This is attained by producing a quadroma fused from hybridomas, one of which produces antibodies that have an affinity to the binding site of protein A and the other of which produces antibodies that have a weaker or no affinity to the binding site of protein A: by propagating and cultivating the quadromas as usual, and by eluting the bsAk in a pH range at least 0.5 units above the pH at which the antibodies with greater affinity to the binding site of protein A are still bonded.

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(54) 【発明の名称】 異種二重特異性抗体の製造法

(57) 【要約】

本発明は、異種二重特異性抗体の製造法に関する。本発明の課題は、高い純度での異種二重特異性抗体を製造することができるような方法を実施することである。前記課題は、1つのハイブリドーマが、蛋白質Aの結合ドメインに対する親和性を有する抗体を生産し、別のハイブリドーマは、第一の抗体と比べて、蛋白質Aの結合ドメインに対して弱い親和性を有するかまたは親和性を有していない抗体を生産するようなハイブリドーマから融合されたクアドロームの製造、酸クアドロームの常法による増殖および培養並びに蛋白質Aの結合ドメインに対してより強い親和性を有する抗体がなお結合しているりHを少なくとも0.5単位上回っているりH範囲内でのbをAkの溶離によって解決される。

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【図1】

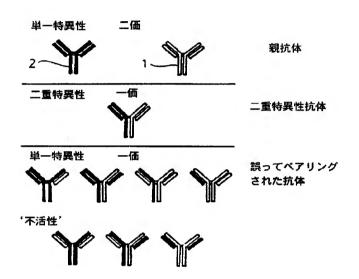
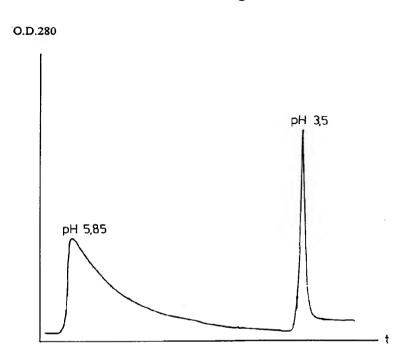
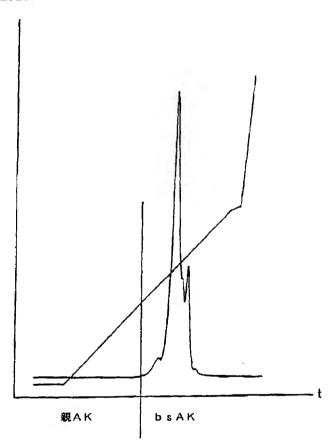


Fig. 2

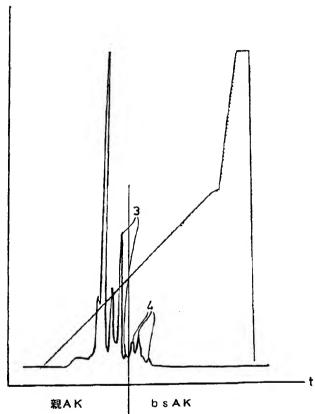












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	SEARCHED		
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	ion searched other than minimum documentation to the extent that		earched
Electronic d	ata dese constilled thiring the international search (name of data be	se and, where practical, search terms used)	
C. DOCUM	ENT'S CONSIDERED TO BE RELEVANT		
Category '	Citation of document, with indication, where appropriate, of the r	elevant passages	Relevant to claim. No.
×	GB.A.2 197 323 (NATIONAL RESEARCE DEVELOPMENT CORPORATION) 18 May 1 see examples 2,6 see claims		1-4
X	GB,A,2 197 322 (NATIONAL RESEARCH DEVELOPMENT CORPORATION) 18 May 1 see examples 5,9 see claims		1
<u> </u>	ner documents are listed in the continuation of box C.	X Patent family members are listed	IR ARINOT.
*Special categories of oited documents: *A" document defitting the general state of the art which is not considered to be of particular relevance. *B" earlier document but published on or after the international filing date. *L" document which may throw doubts on priority distint(s) or which is noted to establish the publication date of another distance or period to the design of the publication date of another distance or period to an oral disclosure, use, chibition or when its noted to establish the publication date of another distance or period to an oral disclosure, use, chibition or when means to be considered to involve an investive step when the document is combined with a publication distance or period to the considered to involve an investive step when the document is combined with a combine			in the application but severy underlying the daimed invention the considered to executed in the con- ection in the con- cept of the con- tent of the con-
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